Appendix A:

Marked-up Copy of Claim Amendments

- 91. (Twice amended) A process of screening a substance for its ability to specifically bind to [interact with] an opioid receptor, said process comprising the steps of:
 - a) expressing a recombinant opioid receptor polypeptide encoded for by a nucleic acid sequence comprising at least 30 contiguous bases of SEQ ID NO:1;
 - b) contacting said substance with the opioid receptor polypeptide; and
 - c) detecting the ability of said substance to <u>specifically bind to</u> [interact with] said opioid receptor polypeptide.
- 97. (Twice amended) A process of screening a substance for its ability to <u>specifically</u> bind to [interact with] an opioid receptor, said process comprising the steps of:
 - expressing a recombinant opioid receptor polypeptide encoded for by a nucleic acid sequence comprising a segment consisting of [at least] 30 contiguous bases of SEQ ID NO:11;
 - b) contacting said substance with the opioid receptor polypeptide; and
 - c) detecting the ability of said substance to <u>specifically bind to</u> [interact with] said opioid receptor polypeptide.
- 98. (Twice amended) The process of claim 97, wherein said opioid receptor polypeptide is encoded for by a nucleic acid sequence comprising a segment consisting of [at least] 40 contiguous bases of SEQ ID NO:11.
- 99. (Twice amended) The process of claim 98, wherein said opioid receptor polypeptide is encoded for by a nucleic acid sequence comprising a segment consisting of [at least] 50 contiguous bases of SEQ ID NO:11.

- 100. (Twice amended) The process of claim 99, wherein said opioid receptor polypeptide is encoded for by a nucleic acid sequence comprising a segment consisting of [at least] 75 contiguous bases of SEQ ID NO:11.
- 101. (Twice amended) The process of claim 100, wherein said opioid receptor polypeptide is encoded for by a nucleic acid sequence comprising a segment consisting of [at least] 100 contiguous bases of SEQ ID NO:11.
- 102. (Twice amended) The process of claim 101, wherein said opioid receptor polypeptide is encoded for by a nucleic acid sequence comprising a segment consisting of [at least] 680 contiguous bases of SEQ ID NO:11.
- 109. (Twice amended) A process of isolating a substance with an ability to act as a specific agonist of a kappa opioid receptor, said process comprising the steps of:
 - a) providing an opioid receptor polypeptide <u>comprising the second</u>

 <u>extracellular loop and</u> encoded for by a nucleic acid sequence comprising

 <u>a segment consisting of 60</u> [at least 30] contiguous bases of SEQ ID

 NO:11;
 - b) contacting said opioid receptor polypeptide with a composition comprising said substance;
 - c) detecting the ability of said substance to interact as an agonist with said opioid receptor polypeptide; and
 - d) isolating said substance if the ability of said substance to <u>specifically bind</u>
 to [interact with] the opioid receptor polypeptide is detected.
- 112. (Twice amended) The process of claim 111, wherein said opioid receptor polypeptide is encoded for by a nucleic acid sequence comprising a segment consisting of [at least] 75 contiguous bases of SEQ ID NO:11.

- 113. (Twice amended) The process of claim 112, wherein said opioid receptor polypeptide is encoded for by a nucleic acid sequence comprising <u>a segment consisting of</u> [at least] 100 contiguous bases of SEQ ID NO:11.
- 114. (Twice amended) The process of claim 113, wherein said opioid receptor polypeptide is encoded for by a nucleic acid sequence comprising a segment consisting of [at least] 680 contiguous bases of SEQ ID NO:11.
- 115. (Amended) A process of screening a substance for its ability to specifically bind to an opioid receptor comprising:
 - a) expressing either (1) a recombinant opioid receptor polypeptide encoded for by a nucleic acid sequence comprising at least 30 contiguous bases of SEQ ID NO:1 or (2) a recombinant opioid receptor polypeptide comprising the second extracellular loop and encoded for by a nucleic acid sequence comprising a segment consisting of 60 [at least 30] contiguous bases of SEQ ID NO:11;
 - b) contacting said substance with the opioid receptor polypeptide; and
 - c) detecting the ability of said substance to interact with said opioid receptor polypeptide.
- 121. (Amended) The process according to claim 115, wherein the opioid receptor polypeptide is a kappa opioid receptor polypeptide having the sequence of SEQ ID NO:2 or comprising a segment consisting of SEQ ID NO:12.
- 124. (Amended) A process of isolating a substance with an ability to act as a agonist of a kappa opioid receptor comprising:
 - a) providing a recombinant opioid receptor polypeptide that includes the second extracellular loop and that is encoded for by a nucleic acid sequence comprising at least 30 contiguous bases of SEQ ID NO:1 or a segment consisting of SEQ ID NO:11;

- b) contacting said opioid receptor polypeptide with a composition comprising the substance;
- c) detecting the ability of the substance to interact as an agonist with the opioid receptor polypeptide; and
- d) isolating the substance if an ability of the substance to interact with the opioid receptor polypeptide is detected.
- 129. (Amended) A process of screening a substance for its ability to act as an agonist of a kappa opioid receptor comprising:
 - a) expressing either (1) a chimeric recombinant opioid receptor polypeptide comprising a portion of the second extracellular loop and encoded for by a nucleic acid sequence comprising at least 30 contiguous bases of SEQ ID NO:1 or (2) a chimeric recombinant opioid receptor polypeptide comprising the second extracellular loop and encoded for by a nucleic acid sequence comprising a segment consisting of 60 [at least 30] contiguous bases of SEQ ID NO:11;
 - b) contacting said substance with the opioid receptor polypeptide; and
 - c) detecting the ability of the substance to interact as an agonist with the opioid receptor polypeptide.

APPENDIX B

PENDING CLAIMS AS OF APRIL 30, 2001

- 91. A process of screening a substance for its ability to specifically bind to an opioid receptor, said process comprising the steps of:
 - a) expressing a recombinant opioid receptor polypeptide encoded for by a nucleic acid sequence comprising at least 30 contiguous bases of SEQ ID NO:1;
 - b) contacting said substance with the opioid receptor polypeptide; and
 - c) detecting the ability of said substance to specifically bind to said opioid receptor polypeptide.
- 92. The process of claim 91, wherein said opioid receptor polypeptide is encoded for by a nucleic acid sequence comprising at least 40 contiguous bases of SEQ ID NO:1.
- 93. The process of claim 92, wherein said opioid receptor polypeptide is encoded for by a nucleic acid sequence comprising at least 50 contiguous bases of SEQ ID NO:1.
- 94. The process of claim 93, wherein said opioid receptor polypeptide is encoded for by a nucleic acid sequence comprising at least 75 contiguous bases of SEQ ID NO:1.
- 95. The process of claim 94, wherein said opioid receptor polypeptide is encoded for by a nucleic acid sequence comprising at least 100 contiguous bases of SEQ ID NO:1.
- 96. The process of claim 95, wherein said opioid receptor polypeptide is encoded for by a nucleic acid sequence comprising at least 680 contiguous bases of SEQ ID NO:1.
- 97. A process of screening a substance for its ability to specifically bind to an opioid receptor, said process comprising the steps of:

- a) expressing a recombinant opioid receptor polypeptide encoded for by a nucleic acid sequence comprising a segment consisting of 30 contiguous bases of SEQ ID NO:11;
- b) contacting said substance with the opioid receptor polypeptide; and
- c) detecting the ability of said substance to specifically bind to said opioid receptor polypeptide.
- 98. The process of claim 97, wherein said opioid receptor polypeptide is encoded for by a nucleic acid sequence comprising a segment consisting of 40 contiguous bases of SEQ ID NO:11.
- 99. The process of claim 98, wherein said opioid receptor polypeptide is encoded for by a nucleic acid sequence comprising a segment consisting of 50 contiguous bases of SEQ ID NO:11.
- 100. The process of claim 99, wherein said opioid receptor polypeptide is encoded for by a nucleic acid sequence comprising a segment consisting of 75 contiguous bases of SEQ ID NO:11.
- 101. The process of claim 100, wherein said opioid receptor polypeptide is encoded for by a nucleic acid sequence comprising a segment consisting of 100 contiguous bases of SEQ ID NO:11.
- 102. The process of claim 101, wherein said opioid receptor polypeptide is encoded for by a nucleic acid sequence comprising a segment consisting of 680 contiguous bases of SEQ ID NO:11.
- 103. A process of isolating a substance with an ability to act as a specific agonist of a kappa opioid receptor, said process comprising the steps of:

- a) providing an opioid receptor polypeptide comprising a second extracellular loop and encoded for by a nucleic acid sequence comprising at least 30 contiguous bases of SEQ ID NO:1;
- b) contacting said opioid receptor polypeptide with a composition comprising said substance;
- c) detecting the ability of said substance to interact as an agonist with said opioid receptor; and
- d) isolating said substance if the ability of said substance to specifically interact with the opioid receptor is detected.
- 104. The process of claim 103, wherein said opioid receptor polypeptide is encoded for by a nucleic acid sequence comprising at least 40 contiguous bases of SEQ ID NO:1.
- 105. The process of claim 104, wherein said opioid receptor polypeptide is encoded for by a nucleic acid sequence comprising at least 50 contiguous bases of SEQ ID NO:1.
- 106. The process of claim 105, wherein said opioid receptor polypeptide is encoded for by a nucleic acid sequence comprising at least 75 contiguous bases of SEQ ID NO:1.
- 107. The process of claim 106, wherein said opioid receptor polypeptide is encoded for by a nucleic acid sequence comprising at least 100 contiguous bases of SEQ ID NO:1.
- 108. The process of claim 107, wherein said opioid receptor polypeptide is encoded for by a nucleic acid sequence comprising at least 680 contiguous bases of SEQ ID NO:1.
- 109. A process of isolating a substance with an ability to act as a specific agonist of a kappa opioid receptor, said process comprising the steps of:

- a) providing an opioid receptor polypeptide comprising the second extracellular loop and encoded for by a nucleic acid sequence comprising a segment consisting of 60 contiguous bases of SEQ ID NO:11;
- b) contacting said opioid receptor polypeptide with a composition comprising said substance;
- c) detecting the ability of said substance to interact as an agonist with said opioid receptor polypeptide; and
- d) isolating said substance if the ability of said substance to specifically bind to the opioid receptor polypeptide is detected.
- 112. The process of claim 111, wherein said opioid receptor polypeptide is encoded for by a nucleic acid sequence comprising a segment consisting of 75 contiguous bases of SEQ ID NO:11.
- 113. The process of claim 112, wherein said opioid receptor polypeptide is encoded for by a nucleic acid sequence comprising a segment consisting of 100 contiguous bases of SEQ ID NO:11.
- 114. The process of claim 113, wherein said opioid receptor polypeptide is encoded for by a nucleic acid sequence comprising a segment consisting of 680 contiguous bases of SEQ ID NO:11.
- 115. A process of screening a substance for its ability to specifically bind to an opioid receptor comprising:
 - a) expressing either (1) a recombinant opioid receptor polypeptide encoded for by a nucleic acid sequence comprising at least 30 contiguous bases of SEQ ID NO:1 or (2) a recombinant opioid receptor polypeptide comprising the second extracellular loop and encoded for by a nucleic acid

sequence comprising a segment consisting of 60 contiguous bases of SEQ ID NO:11;

- b) contacting said substance with the opioid receptor polypeptide; and
- detecting the ability of said substance to interact with said opioid receptor polypeptide.
- 116. The process according to claim 115, wherein said opioid receptor polypeptide is a chimeric opioid receptor polypeptide.
- 117. The process of claim 116, wherein one polypeptide of the chimeric opioid receptor polypeptide comprises the second extracellular loop of kappa opioid receptor.
- 118. The process of claim 116, wherein one polypeptide of the chimeric opioid receptor polypeptide comprises the third extracellular loop of kappa opioid receptor.
- 119. The process of claim 116, wherein the chimeric opioid receptor polypeptide comprises polypeptide portions of both kappa and delta opioid receptors.
- 120. The process according to claim 116, wherein the chimeric opioid receptor polypeptide comprises 1-78/70-372 or 1-69/79-380.
- 121. The process according to claim 115, wherein the opioid receptor polypeptide is a kappa opioid receptor polypeptide having the sequence of SEQ ID NO:2 or comprising a segment consisting of SEQ ID NO:12.
- 122. The process of claim 121, wherein said opioid receptor polypeptide is a kappa opioid receptor polypeptide encoded for by the polynucleotide of SEQ ID NO: 1.

- 123. The process of claim 121, wherein said opioid receptor polypeptide is a kappa opioid receptor polypeptide encoded for by the polynucleotide of SEQ ID NO: 11.
- 124. (Amended) A process of isolating a substance with an ability to act as a agonist of a kappa opioid receptor comprising:
 - a) providing a recombinant opioid receptor polypeptide that includes the second extracellular loop and that is encoded for by a nucleic acid sequence comprising at least 30 contiguous bases of SEQ ID NO:1 or a segment consisting of SEQ ID NO:11;
 - b) contacting said opioid receptor polypeptide with a composition comprising the substance;
 - c) detecting the ability of the substance to interact as an agonist with the opioid receptor polypeptide; and
 - d) isolating the substance if an ability of the substance to interact with the opioid receptor polypeptide is detected.
- 125. The process of claim 124, wherein the opioid receptor polypeptide is a chimeric opioid receptor polypeptide.
- 126. The process of claim 124, wherein one polypeptide of the chimeric opioid receptor polypeptide comprises the third extracellular loop of delta opioid receptor.
- 127. The process of claim 124, wherein the opioid receptor polypeptide comprises portions of both kappa and delta opioid receptors.
- 128. The process of claim 124, wherein the chimeric polypeptide comprises 1-78/70-372 or 1-69/79-380.

- 129. A process of screening a substance for its ability to act as an agonist of a kappa opioid receptor comprising:
 - a) expressing either (1) a chimeric recombinant opioid receptor polypeptide comprising a portion of the second extracellular loop and encoded for by a nucleic acid sequence comprising at least 30 contiguous bases of SEQ ID NO:1 or (2) a chimeric recombinant opioid receptor polypeptide comprising the second extracellular loop and encoded for by a nucleic acid sequence comprising a segment consisting of 60 contiguous bases of SEQ ID NO:11;
 - b) contacting said substance with the opioid receptor polypeptide; and
 - c) detecting the ability of the substance to interact as an agonist with the opioid receptor polypeptide.
- 130. The process of claim 129, wherein said nucleic acid sequence comprises at least 40 contiguous bases of SEQ ID NO:1.
- 131. The process of claim 129, wherein said nucleic acid sequence comprises at least 55 contiguous bases of SEQ ID NO:1.
- 132. The process of claim 129, wherein said nucleic acid sequence comprises at least 70 contiguous bases of SEQ ID NO:1.
- 133. The process of claim 129, wherein one polypeptide of the chimeric opioid receptor polypeptide comprises the second extracellular loop of kappa opioid receptor.
- 134. The process of claim 129, wherein one polypeptide of the chimeric opioid receptor polypeptide comprises the third extracellular loop of kappa opioid receptor.

- 135. The process of claim 129, wherein the chimeric opioid receptor polypeptide comprises polypeptide portions of both kappa and delta opioid receptors.
- 136. The process of claim 97, wherein the recombinant opioid receptor polypeptide comprises the second extracellular loop.